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APPLICATION NO.	FILI	NG DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.	
09/485,071	71 02/03/2000		NORBERT O. REICH	30794.30USWO	1444	
22462	7590	06/02/2005		EXAMINER		
GATES &			LEWIS, PATRICK T			
HOWARD HUGHES CENTER 6701 CENTER DRIVE WEST, SUITE 1050				ART UNIT	PAPER NUMBER	
LOS ANGELES, CA 90045				1623	1623	

DATE MAILED: 06/02/2005

Please find below and/or attached an Office communication concerning this application or proceeding.

## Advisory Action Before the Filing of an Appeal Brief

Application No.	Applicant(s)	
09/485,071	REICH ET AL.	
Examiner	Art Unit	
Patrick T. Lewis	1623	

--The MAILING DATE of this communication appears on the cover sheet with the correspondence address --THE REPLY FILED 09 May 2005 FAILS TO PLACE THIS APPLICATION IN CONDITION FOR ALLOWANCE. 1. 🔀 The reply was filed after a final rejection, but prior to or on the same day as filing a Notice of Appeal. To avoid abandonment of this application, applicant must timely file one of the following replies: (1) an amendment, affidavit, or other evidence, which places the application in condition for allowance; (2) a Notice of Appeal (with appeal fee) in compliance with 37 CFR 41.31; or (3) a Request for Continued Examination (RCE) in compliance with 37 CFR 1.114. The reply must be filed within one of the following time periods: The period for reply expires \_\_\_\_ months from the mailing date of the final rejection. b) X The period for reply expires on: (1) the mailing date of this Advisory Action, or (2) the date set forth in the final rejection, whichever is later. In no event, however, will the statutory period for reply expire later than SIX MONTHS from the mailing date of the final rejection. Examiner Note: If box 1 is checked, check either box (a) or (b), ONLY CHECK BOX (b) WHEN THE FIRST REPLY WAS FILED WITHIN TWO MONTHS OF THE FINAL REJECTION. See MPEP 706.07(f). Extensions of time may be obtained under 37 CFR 1.136(a). The date on which the petition under 37 CFR 1.136(a) and the appropriate extension fee have been filed is the date for purposes of determining the period of extension and the corresponding amount of the fee. The appropriate extension fee under 37 CFR 1.17(a) is calculated from: (1) the expiration date of the shortened statutory period for reply originally set in the final Office action; or (2) as set forth in (b) above, if checked. Any reply received by the Office later than three months after the mailing date of the final rejection, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b). **NOTICE OF APPEAL** 2. The Notice of Appeal was filed on \_\_\_\_\_. A brief in compliance with 37 CFR 41.37 must be filed within two months of the date of filing the Notice of Appeal (37 CFR 41.37(a)), or any extension thereof (37 CFR 41.37(e)), to avoid dismissal of the appeal. Since a Notice of Appeal has been filed, any reply must be filed within the time period set forth in 37 CFR 41.37(a). **AMENDMENTS** 3. The proposed amendment(s) filed after a final rejection, but prior to the date of filing a brief, will not be entered because (a) They raise new issues that would require further consideration and/or search (see NOTE below); (b) They raise the issue of new matter (see NOTE below); (c) They are not deemed to place the application in better form for appeal by materially reducing or simplifying the issues for appeal; and/or (d) They present additional claims without canceling a corresponding number of finally rejected claims. NOTE: . (See 37 CFR 1.116 and 41.33(a)). 4. The amendments are not in compliance with 37 CFR 1.121. See attached Notice of Non-Compliant Amendment (PTOL-324). 5. Applicant's reply has overcome the following rejection(s): \_\_\_ 6. Newly proposed or amended claim(s) \_\_\_\_\_ would be allowable if submitted in a separate, timely filed amendment canceling the non-allowable claim(s). 7. To purposes of appeal, the proposed amendment(s): a) will not be entered, or b) will be entered and an explanation of how the new or amended claims would be rejected is provided below or appended. The status of the claim(s) is (or will be) as follows: Claim(s) allowed: Claim(s) objected to: Claim(s) rejected: \_\_ Claim(s) withdrawn from consideration: AFFIDAVIT OR OTHER EVIDENCE 8. The affidavit or other evidence filed after a final action, but before or on the date of filing a Notice of Appeal will not be entered because applicant failed to provide a showing of good and sufficient reasons why the affidavit or other evidence is necessary and was not earlier presented. See 37 CFR 1.116(e). 9. The affidavit or other evidence filed after the date of filing a Notice of Appeal, but prior to the date of filing a brief, will not be entered because the affidavit or other evidence failed to overcome all rejections under appeal and/or appellant fails to provide a showing a good and sufficient reasons why it is necessary and was not earlier presented. See 37 CFR 41.33(d)(1). 10. The affidavit or other evidence is entered. An explanation of the status of the claims after entry is below or attached. REQUEST FOR RECONSIDERATION/OTHER 11. \times The request for reconsideration has been considered but does NOT place the application in condition for allowance because: See Continuation Sheet. 12. Note the attached Information Disclosure Statement(s). (PTO/SB/08 or PTO-1449) Paper No(s). 13. Other: \_\_\_\_\_.

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PTOL-303 (Rev. 4-05)

Continuation of 11. does NOT place the application in condition for allowance because: Flynn teaches substrates designed to mimic DNA transcriptional cis elements previously reported to have cytosine C-5-methylated regulation. The primary focus on the Flynn study was on the two DNA sequences shown in Table 1 (GC-box bmet reads upon a synthetic oligonucleotide of 30 nucleotides in length which comprises a 5mCpG dinucleotide and the nucleotide sequence shown in SEQ ID NO: 10). The two sequences are thought to differentially regulate gene expression, depending on the methylation state of a single CpG dinucleotide. The rate-limiting step for these substrates is the methylation step itself. Methylation of DNA at cytosine (5-mC)1 occurs in most biological kingdoms. The function of cytosine methylation in mammals mainly involves positive and negative transcriptional control and appears predominantly in the minimal context of the CpG dinucleotide. Amplification of DCMTase gene induces tumorigenic transformation of NIH 3T3 mouse fibroblasts; correspondingly, human neoplastic cells and cells derived from different stages of colon cancer expresses up to 200-fold higher levels of DCMTase than normal. Conversely, expression of antisense DCMTase mRNA is the adrenocortical carcinoma cell line Y1 inhibits tumorigenesis. The anticancer agent 5-azadeoxycytidine functions by inhibiting the DCMTase, and DCMTase activity contributes substantially to tumor development in a mouse model of intestinal neoplasia. Flynn further explains on pages 7309-7310 that a precise functional description of the enzyme is essential for understanding how DCMTase methylation and for the design of novel anticancer strategies based on regulation of the enzyme. Flynn's explanation is seen to be sufficient motivation to modify the internucleotide linkages and provide a pharmaceutical composition as instantly claimed.